# Infections in Combat Casualties During Operations Iraqi and Enduring Freedom

Clinton K. Murray, MD, Kenneth Wilkins, PhD, Nancy C. Molter, PhD, Heather C. Yun, MD, Michael A. Dubick, PhD, Mary Ann Spott, MPA, MBA, Donald Jenkins, MD, Brian Eastridge, MD, John B. Holcomb, MD, Lorne H. Blackbourne, MD, and Duane R. Hospenthal, MD, PhD

**Background:** Infections are a common acute and chronic complication of combatrelated injuries; however, no systematic attempt to assess infections associated with US combat-related injuries occurring in Iraq and Afghanistan has been conducted. The Joint Theater Trauma Registry (JTTR) has been established to collect injury specific medical data from casualties in Iraq and Afghanistan.

**Methods:** We reviewed the JTTR for the identification of infectious complications (IC) using International Classification of Diseases, 9th Revision (ICD-9) coding during two phases of the wars, before and after the end of the major ground

operations in Iraq (19 March-May 31, 2003 and June 1, 2003-December 31, 2006). ICD-9 codes were combined into two categories; anatomic or clinical syndrome and pathogen. An IC was defined as the presence of ICD-9 codes that included both anatomic or clinical syndrome and a pathogen.

**Results:** There were 425 patients evaluated in phase I and 684 in phase II with approximately one third having an IC. The most common anatomic or clinical syndrome codes were skin or wound followed by lung, and the most common pathogen code was gram-negative bacteria. The site of injury had varying rates of

IC: spine or back (53%), head or neck (44%), torso (43%), and extremity (35%). Injury Severity Score and certain mechanisms of injury (explosive device, bomb, and landmine) were associated with an IC on multivariate analysis (p < 0.01).

**Conclusion:** Infections are common after combat-related injuries. Although the JTTR can provide general information regarding infections, improved data capture and more specific clinical information is necessary to improve overall combat-related injury infection care.

**Key Words:** Combat, Infection, Iraq, Afghanistan.

J Trauma. 2009;66:S138-S144.

he deployment of forward surgical assets, rapid evacuation to medical care, use of body armor, and well-trained corpsmen and medics have culminated in improved care and outcomes of US combat casualties. One of the priorities of combat casualty care is to continually assess ongoing management strategies to improve care. The Joint Theater Trauma Registry (JTTR) was established as a performance improvement program to provide a central repository of clinical data from all casualties injured in Operation Iraqi and

Submitted for publication January 22, 2009. Accepted for publication January 22, 2009.

Copyright © 2009 by Lippincott Williams & Wilkins

From the Brooke Army Medical Center (C.K.M., H.C.Y., B.E., J.B.H., L.H.B., D.R.H.), Fort Sam Houston, Texas; Department of Preventive Medicine and Biometrics (C.K.M., K.W., H.C.Y., D.R.H.) Uniformed Services University of the Health Sciences, Bethesda, Maryland; Infectious Disease Clinical Research Program (K.W.), Bethesda, Maryland; and US Army Institute of Surgical Research (N.C.M., M.A.D., M.A.S., D.J., B.E., J.B.H., L.H.B.), Fort Sam Houston, Texas.

The opinions or assertions contained herein are the private views of the authors and are not to be construed as official or reflecting the views of the Department of the Army, Department of the Air Force, Department of Defense or the US government. This work was prepared as part of their official duties and, as such, there is no copyright to be transferred.

Address for reprints: Clinton K. Murray, MD, Infectious Disease Service, San Antonio Military Medical Center, Brooke Army Medical Center, 3851 Roger Brooke Drive, Fort Sam Houston, TX 78234; email: Clinton. Murray@amedd.army.mil.

DOI: 10.1097/TA.0b013e31819d894c

Enduring Freedom (OIF or OEF) to allow assessment of injury patterns, care provided, and associated outcomes.<sup>1</sup>

One of the lessons learned during previous wars is that infections associated with combat-related injuries can have significant impact on morbidity and mortality. Septic shock was a leading cause of mortality in the Vietnam War with infections noted to complicate 3.9% of casualties.<sup>2–4</sup> During the current wars in Iraq and Afghanistan, infections are also noted to complicate the care of combat casualties. An autopsy review of US casualties of OIF or OEF during two separate time periods of the war revealed sepsis as the fourth leading cause of potentially survivable injury, behind central nervous system injury, hemorrhage, and airway disorders.<sup>5</sup> Among extremity injuries in US casualties, approximately 5% to 15% develop osteomyelitis.<sup>6-9</sup> One of the major concerns with combat-related infections is that they are associated with bacteria that are multidrug resistant, limiting potential therapy.<sup>7-9</sup> However, no study has been undertaken to evaluate large numbers of casualties for injury patterns and associated infectious complications (IC). This study used the JTTR to capture data from all completed records for the presence of infectious codes among personnel with combat-related injuries associated with service in OIF or OEF.

## **MATERIALS AND METHODS**

The JTTR is a Department of Defense (DoD) performance improvement program designed to collect medical data from patients that suffer trauma within the current com-

S138 April Supplement 2009

| maintaining the data needed, and c<br>including suggestions for reducing | completing and reviewing the collect<br>this burden, to Washington Headqu<br>uld be aware that notwithstanding an | o average 1 hour per response, includion of information. Send comments a arters Services, Directorate for Informy other provision of law, no person to | regarding this burden estimate mation Operations and Reports | or any other aspect of th<br>, 1215 Jefferson Davis l | is collection of information,<br>Highway, Suite 1204, Arlington |
|--|---|--|--|---|---|
| 1. REPORT DATE<br>01 APR 2009  |   | 2. REPORT TYPE <b>N/A</b>  |  | 3. DATES COVE   | RED   |
| 4. TITLE AND SUBTITLE  |   |  |  | 5a. CONTRACT  | NUMBER  |
| Infections in comba  | at casualties during  | Operations Iraqi an  | d Enduring   | 5b. GRANT NUM   | 1BER  |
| rreedom  |   |  |  | 5c. PROGRAM E   | LEMENT NUMBER   |
| 6. AUTHOR(S)   |   |  |  | 5d. PROJECT NU  | MBER  |
| • /  | *   | C., Yun H. C., Dubic<br>mb J. B., Blackbour  | ′ <del>-</del>   | 5e. TASK NUMB   | ER  |
| Hospenthal D. R.,  |   |  |  | 5f. WORK UNIT   | NUMBER  |
|  | •   | odress(es)<br>al Research, JBSA l  | Fort Sam   | 8. PERFORMING<br>REPORT NUMBI                         | GORGANIZATION<br>ER   |
| 9. SPONSORING/MONITO   | RING AGENCY NAME(S) A   | AND ADDRESS(ES)  |  | 10. SPONSOR/M   | ONITOR'S ACRONYM(S)   |
|  |   |  |  | 11. SPONSOR/M<br>NUMBER(S)                            | ONITOR'S REPORT   |
| 12. DISTRIBUTION/AVAIL Approved for publ                                 | LABILITY STATEMENT<br>ic release, distributi  | on unlimited   |  |   |   |
| 13. SUPPLEMENTARY NO   | OTES  |  |  |   |   |
| 14. ABSTRACT   |   |  |  |   |   |
| 15. SUBJECT TERMS  |   |  |  |   |   |
| 16. SECURITY CLASSIFIC   | CATION OF:  |  | 17. LIMITATION OF<br>ABSTRACT                                | 18. NUMBER  | 19a. NAME OF<br>RESPONSIBLE PERSON                              |
| a. REPORT<br>unclassified  | b. ABSTRACT <b>unclassified</b>   | c. THIS PAGE unclassified  | UU   | OF PAGES<br><b>7</b>                                  | RESTUNSIBLE PERSON  |

**Report Documentation Page** 

Form Approved OMB No. 0704-0188 bat theaters in Iraq and Afghanistan. The JTTR contains data that is collected and summarized in a central repository, from point of injury through the various levels of medical care in and out of the combat zone. This study was reviewed and approved by the Brooke Army Medical Center Institutional Review Board and the Unites States Army Institute of Surgical Research.

This study assessed the JTTR for the presence of infections associated with injuries sustained during combat operations in Iraq and Afghanistan. Data recovery from the JTTR for use in this study was completed on January 2, 2008. The study evaluated two phases of the wars: phase I from March 19, 2003 to May 31, 2003 defined as the time period from the onset until it was announced that major military operations had been completed; and phase II which was June 1, 2003 to December 31, 2006. Casualties from OEF were also divided into these two time periods. The number of JTTR entries that underwent complete coding was compared with the Patient Administration Systems and Biostatistics Activities (PASBA), which received all inpatient records from deployed medical units.

The data collected included summary totals of patients with data captured in the JTTR by year, theater of operation, and phase of military operations, with associated gender, military rank, branch of military service, injury severity score (ISS), medical care level at the time of diagnosis (level I, II, III, IV, and V), 10 battle injury or nonbattle injury, injury site(s), and mechanism(s) of injury. Injury patterns were grouped by anatomic distribution; specifically, recorded International Classification of Diseases, 9th Revision (ICD-9) codes involving trauma were mapped to body region using the Barell Matrix posted by the CDC (http://www.cdc.gov/ nchs/data/ice/final\_matrix\_post\_ice.pdf). ICs were captured using ICD-9 codes entered within the JTTR. In addition, they were bundled for analysis based upon major categories of anatomic or clinical syndromes, which included bacteremia, abdominal, bone or joint, skin, lung, urine, central nervous system, and other. Infectious disease ICD-9 codes were also bundled into gram-positive bacteria, gram-negative bacteria, other bacteria, and fungi. An IC was defined as a patient having codes for both an anatomic or clinical syndrome and a corresponding pathogen. Note that certain ICD-9 codes for anatomic or clinical syndrome included a pathogen, whereas those ICD-9 codes for anatomic or clinical syndrome lacking a pathogen were considered an IC if a separate ICD-9 code for that patient also specified a pathogen.

Statistical analysis included descriptive evaluation of various cohorts, point and interval estimates for standard logistic and "exact" logistic regression coefficients within multivariate analyses, whereas univariate or bivariate analyses included the unpaired Student's t test or Mann-Whitney rank-sum test for continuous variables, and  $\chi^2$  or Fisher's exact for categorical variables, where appropriate. A nominal 0.05 significance level was employed throughout these analyses.

**Table 1** Total Number of Individual Patients Records Reviewed From the JTTR and Recorded Infectious Disease ICD-9 Codes Associated With Them

| Combat Zone        | Number of<br>Patients | Number of ICD-9 Codes by Year of<br>Trauma |       |       |       |      |  |  |
|--------------------|-----------------------|--|-------|-------|-------|------|--|--|
|                    | i attents             | Total                                      | 2003  | 2004  | 2005  | 2006 |  |  |
| OIF/OEF            | 1,108                 | 12,410                                     | 4,713 | 3,675 | 3,694 | 328  |  |  |
| OIF                | 1,059                 | 11,921                                     | 4,481 | 3,641 | 3,506 | 293  |  |  |
| Battle injuries    | 653                   | 9,085                                      | 2,551 | 3,154 | 3,133 | 247  |  |  |
| Nonbattle injuries | 406                   | 2,836                                      | 1,930 | 487   | 373   | 46   |  |  |
| OEF                | 49                    | 489  | 232   | 34    | 188   | 35   |  |  |
| Battle injuries    | 27                    | 333  | 162   | 29    | 128   | 14   |  |  |
| Nonbattle injuries | 22                    | 156  | 70    | 5     | 60    | 21   |  |  |

### **RESULTS**

There were completed records for 562 individuals injured in 2003, 284 for 2004, 231 for 2005, and 31 for 2006, representing a 100%, 77%, 66%, and 23% completion rates based on PASBA data, respectively (Table 1). Overall, only 56% of records had been entered into JTTR as of November 2007 based on PASBA data.

Nearly, all of the records reviewed were from patients injured in Iraq (96%) (Tables 1 and 2). Overall, 61% of injuries were due to battle injuries although there were fewer battle injuries in phase I (38%) compared with phase II (76%) of the war (Table 2). Most casualties were Army personnel although a higher percentage of Marines was noted in phase I. The majority of patients were enlisted (grade of E-5 or below; mostly E-4) men. Explosion associated injuries were more common during phase II (80%) than during phase I (42%), with grenade associated explosions being the most common in phase I (24%) and improvised explosions being the most common in phase II (32%) than phase II (16%) among battle related injuries. Casualties in phase II (15) had greater mean ISS scores than during phase I (4).

During phase I only 28 of the 425 patients (6.6%) had an ICD-9 code for an infectious anatomic or clinical syndrome or pathogen in contrast to phase II, where 661 of 684 patients (96.6%) had an ICD-9 code for an infectious anatomic or clinical syndrome or pathogen. The most common infectious code by anatomic or clinical syndrome was skin and wound followed by lung (Table 3). Gram-negative bacteria were the most common pathogens reported (Table 3). Overall, an IC was present in 35% of patients in phase II. The specific ICD-9 codes for anatomic or clinical syndrome or pathogen are provided in Tables 4 and 5, respectively, along with the frequency reported for each code in the abstracted JTTR data. Based on anatomic site of injury, 55% of spine and back injuries, 44% of torso injuries, 43% of head and neck injuries, and 34% of extremity injuries developed an IC during phase II (Table 6). The first level of military medical care of an IC was recorded at was level V in 71% of the cases followed by level IV in approximately 27% of cases (Table 7).

*Volume 66* • *Number 4* **\$139** 

**Table 2** Patient Demographics in the JTTR for Phase I (19 March 2003–30 May 2003) and Phase II (1 June 2003–31 December 2006)

|                                  | Phase I   | Phase II                                     |            |       |      |      |      |      |       |     |  |
|----------------------------------|---|--|------------|-------|------|------|------|------|-------|-----|--|
|                                  | Number (%)  | Total, Number (%)                            | 2003       | 200   | 04   |      | 2005 |      | 20    | 06  |  |
| Number of patients               | 425   | 684  | 137        | 284   | 1    |      | 231  |      | 31    |     |  |
| Female*                          | 21 (4.9%)   | 23 (3.4%)                                    | 4          | 13    | 3    |      | 4    |      | 2     |     |  |
| Branch of service                |   |  |            |       |      |      |      |      |       |     |  |
| Army                             | 319 (75.1%)   | 577 (84.4%)                                  | 134        | 248   | 3    |      | 174  |      | 20    |     |  |
| Marine                           | 88 (20.7%)  | 86 (12.6%)                                   | 1          | 26    | 3    |      | 49   |      | 10    |     |  |
| Air force                        | 7 (1.6%)  | 9 (1.3%)                                     | 1          | 5     | 5    |      | 3    |      | 0     |     |  |
| Navy                             | 11 (2.6%)   | 12 (1.8%)                                    | 1          | 5     | 5    |      | 5    |      | 1     |     |  |
| Military grade/rank              |   |  |            |       |      |      |      |      |       |     |  |
| Enlisted                         | 384 (90.3%)   | 637 (93.3%)                                  | 128        | 264   | 1    |      | 218  |      | 27    |     |  |
| E1                               | 9 (2.1%)  | 6 (0.9%)                                     | 2          | (     | )    |      | 3    |      | 1     |     |  |
| E2                               | 32 (7.6%)   | 23 (3.4%)                                    | 4          | 8     | 3    |      | 11   |      | 0     |     |  |
| E3                               | 82 (19.3%)  | 122 (17.8%)                                  | 20         | 45    |      |      | 47   |      | 10    |     |  |
| E4                               | 132 (31.1%)   | 241 (35.2%)                                  | 59         | 99    |      |      | 76   |      | 7     |     |  |
| E5                               | 83 (19.5%)  | 134 (19.6%)                                  | 16         | 64    |      |      | 47   |      | 7     |     |  |
| E6                               | 32 (7.5%)   | 70 (10.2%)                                   | 17         | 33    |      |      | 18   |      | 2     |     |  |
| E7                               | 11 (2.6%)   | 31 (4.5%)                                    | 10         | 12    |      |      | 9    |      | 0     |     |  |
| E8                               | ,   | ` ,  | 0          | 3     |      |      | 7    |      | 0     |     |  |
| Warrant                          | 3 (0.7%)  | 10 (1.5%)                                    | 1          | (     |      |      | 0    |      | 1     |     |  |
|                                  | 4 (0.9%)  | 2 (0.3%)                                     |            |       |      |      |      |      |       |     |  |
| W1                               | 2 (0.5%)  | 0 (0.0%)                                     | 0          | (     |      |      | 0    |      | 0     |     |  |
| W2                               | 2 (0.5%)  | 2 (0.3%)                                     | 1          | (     |      |      | 0    |      | 1     |     |  |
| Officer                          | 33 (7.8%)   | 42 (6.1%)                                    | 7          | 20    |      |      | 12   |      | 3     |     |  |
| 01                               | 1 (0.2%)  | 7 (1.0%)                                     | 1          | 3     |      |      | 2    |      | 0     |     |  |
| O2                               | 9 (2.1%)  | 15 (2.2%)                                    | 2          | 3     |      |      | 5    |      | 0     |     |  |
| O3                               | 15 (3.5%)   | 12 (1.8%)                                    | 4          | 5     |      |      | 1    |      | 2     |     |  |
| O4                               | 4 (0.9%)  | 5 (0.7%)                                     | 0          | 3     | 3    |      | 1    |      | 1     |     |  |
| O5                               | 1 (0.2%)  | 1 (0.2%)                                     | 0          | (     | )    |      | 1    |      | 0     |     |  |
| O6                               | 3 (0.7%)  | 3 (0.4%)                                     | 0          | 1     |      |      | 2    |      | 0     |     |  |
| Unknown                          | 4 (0.9%)  | 2 (0.3%)                                     | 1          | (     | )    |      | 1    |      | 0     |     |  |
| Disposition                      |   |  |            |       |      |      |      |      |       |     |  |
| Died                             | 8 (1.9%)  | 11 (1.6%)                                    | 1          | 4     | 1    |      | 4    |      | 2     |     |  |
| Return to duty                   | 55 (12.9%)  | 31 (5.0%)                                    | 10         | 16    | 3    |      | 5    |      | 0     |     |  |
| Unknown                          | 362 (85.2%)   | 642 (93.9%)                                  | 127        | 268   |      |      | 225  |      | 31    |     |  |
| Total-OIF                        | 404 (95.1%)   | 656 (95.9%)                                  | 133        | 279   |      |      | 218  |      | 25    |     |  |
| Battle injury                    | 150 (37.1% of 404)                                  | 504 (76.8% of 656)                           | 87         | 224   |      |      | 175  |      | 17    |     |  |
| Total-OEF                        | 21 (4.9%)   | 28 (4.1%)                                    | 4          | 5     |      |      | 13   |      | 6     |     |  |
| Battle injury                    | 11 (52.4% of 21)                                    | 16 (57.1% of 28)                             | 2          | 2     |      |      | 8    |      | 2     |     |  |
| Injury severity score: mean      | 11 (32.470 01 21)                                   | 10 (37.170 01 20)                            | (n = 126)  | (n =  |      | (n   | = 21 | 7)   | (n =  |     |  |
|                                  |   |  | (11 – 120) | (11 — | 239) | (11  | - 21 | 1)   | (11 — | 21) |  |
| of maximum                       |   |  |            |       |      |      |      |      |       |     |  |
| score (range)                    | F 0 (1 75)  | 00.0 (4. 75)                                 | 10.0       | 0.0   |      |      | 01.0 |      | 47    |     |  |
| Military AIS (n = $630$ )        | 5.2 (1, 75)   | 20.3 (1, 75)                                 | 19.2       |       | ).5  |      | 21.0 |      | 17    |     |  |
| 2005 (n = 630)                   | 4.3 (1, 38)   | 14.6 (1, 75)                                 | 13.3       | 14    | 1.5  |      | 15.8 |      | 13    | .7  |  |
| Mechanism of injury <sup>†</sup> | BI: 161/NBI: 266 (% of 161/<br>of 266) <sup>†</sup> | % Bls: 523/NBls: 165 (% of 165) <sup>†</sup> | % of 523/% |       |      |      |      |      |       |     |  |
| Cymlonian                        |   |  | 2 60/) DI  | NDI   | DI   | NIDI | DI   | NIDI | DI    | ND  |  |
| Explosion                        | 67/13 (41.6%/4.9%)                                  | 419/6 (80.1%/3                               | ,          | NBI   | BI   | NBI  | BI   | NBI  | BI    | NB  |  |
| Gun shot                         | 52/12 (32.3%/4.5%)                                  | 83/14 (15.9%/8                               | ,          | 7     | 34   |      | 28   | 6    | 7     |     |  |
| Shrapnel/fragment                | 17/0 (10.6%/0%)                                     | 3/1 (0.6%/0.                                 | ,          | _     | _    |      | 1    |      |       |     |  |
| Motor vehicle crash              | 3/64 (1.9%/24.1%)                                   | 2/30 (0.4%/18                                | ,          | 6     | 1    | 11   | 1    | 11   |       | 2   |  |
| Fall/jump from height            | 2/35 (1.2%/13.2%)                                   | 1/14 (0.2%/8.                                | ,          | 5     |      | 4    |      | 5    |       |     |  |
| Machinery/equipment              | 3/33 (1.9%/12.4%)                                   | 0/15 (0%/9.19                                | ,          | 4     |      | 5    |      | 3    |       | (   |  |
| Exertion/overexertion            | 2/22 (1.2%/8.3%)                                    | 0/0 (0%/0%)                                  |            |       |      |      |      |      |       |     |  |
| Burn                             | 4/13 (2.5%/4.9%)                                    | 4/10 (0.8%/6.                                | 1%)        | 3     | 2    | 4    | 2    | 2    |       |     |  |
| Knife or other sharp object      | 0/3 (0%/1.1%)                                       | 0/9 (0%/5.59                                 | %)         | 3     |      | 1    |      | 3    |       | 2   |  |
| "Other"                          | 11/71 (6.8%/26.7%)                                  | 11/66 (2.1%/40                               | 0%) 5      | 16    | 4    | 31   | 2    | 16   | 5     | 16  |  |

<sup>\*</sup> One casualty wounded in 2005 has gender "Undetermined."

S140 April Supplement 2009

<sup>&</sup>lt;sup>†</sup> Entries reflect number and percentage of reported mechanisms, not individual patients, who might provide more than one mechanism of injury.

BI, number of battle injuries; NBI, number of nonbattle injuries; RPG, rocket-propelled grenade.

**Table 3** Infection ICD-9 Coding Based Upon Anatomical/Clinical Syndrome or Pathogen Bundled

|                         |         | Phase II by Year |      |      |      |      |  |  |
|-------------------------|---------|------------------|------|------|------|------|--|--|
|                         | Phase I | Total            | 2003 | 2004 | 2005 | 2006 |  |  |
| Anatomical and clinical |         |                  |      |      |      |      |  |  |
| syndrome                |         |                  |      |      |      |      |  |  |
| Abdomen                 | 0       | 13               | 1    | 9    | 1    | 2    |  |  |
| Bacteremia              | 2       | 90               | 20   | 38   | 32   | 0    |  |  |
| Bone and joint          | 1       | 37               | 7    | 17   | 13   | 0    |  |  |
| Central nervous         | 0       | 12               | 5    | 5    | 2    | 0    |  |  |
| system                  |         |                  |      |      |      |      |  |  |
| Lung                    | 6       | 94               | 17   | 41   | 30   | 6    |  |  |
| Skin and wound          | 14      | 190              | 36   | 86   | 56   | 12   |  |  |
| Other                   | 3       | 144              | 36   | 65   | 38   | 5    |  |  |
| Pathogen                |         |                  |      |      |      |      |  |  |
| Gram-positive bacteria  | 3       | 123              | 29   | 57   | 34   | 3    |  |  |
| Gram-negative           | 7       | 299              | 57   | 127  | 109  | 6    |  |  |
| bacteria                |         |                  |      |      |      |      |  |  |
| Fungus                  | 0       | 30               | 6    | 11   | 10   | 3    |  |  |
| Other                   | 4       | 185              | 45   | 72   | 64   | 4    |  |  |
| Infectious complication | 6       | 241              | 55   | 101  | 80   | 5    |  |  |

Patients may have received more than one infectious code.

Risk factors associated with IC include certain injury patterns and mechanisms of injury (Table 8). In addition, patients with higher ISS had higher risk of developing an IC. On multivariate analysis, higher ISS, injury during phase II of the study, and certain mechanisms of injury (explosive device, bomb, and landmine) remained significant.

There were a total of 19 deaths reported in this data set. During phase I, 8 persons died from various injuries: grenade, (1), gunshot (1), motor vehicle crash (1), and unknown (5). During phase II, there were 11 deaths (1 in 2003 from a motor vehicle crash; 4 in 2004 from motor vehicle crash [1], gunshot [1], explosion [1] and burn [1]); 4 in 2005 from explosion (3) and grenade (1); 2 in 2006 from gunshot (1) and explosion (1). None of the phase I deaths had any infectious anatomic or clinical syndrome or pathogens listed as a complication whereas 10 of the 11 phase II deaths were associated with an infection by ICD-9 coding. The presence of one or more IC was not estimated (via exact logistic regression) as being significantly associated with mortality, after adjusting for phase and ISS.

#### DISCUSSION

Military personnel who suffer combat-related injuries are at risk of developing acute and chronic ICs. This is even more concerning as many of these infections are due to multidrugresistant bacteria. Although limited single-site studies of OIF or OEF casualties have assessed specific infections of combat-related injuries, this study uses a more global approach through evaluating the JTTR. Overall, approximately one third of the casualties assessed in this study had the presence of an IC. These infections involved primarily skin and wound sites or the lung, with a high rate of bacteremia. As in most combat studies, the primary bacteria were

**Table 4** Infection ICD-9 Coding by Anatomical/Clinical Syndrome Based

| Syndrome   | Code   | Code Description                  | Number |
|------------|--------|-----------------------------------|--------|
| Abdomen    | 567.2  | Suppurative peritonitis           | 3      |
|            | 567.8  | Peritonitis                       | 2      |
|            | 567.9  | Peritonitis unspecified           | 16     |
|            | 569.61 | Colostomy/enterostomy infection   | 3      |
| Bacteremia | 038.11 | Staphylococcus aureus septicemia  | 2      |
|            | 038.43 |                                   | 3      |
|            | 038.49 | Gram-negative septicemia          | 2      |
|            | 038.8  | Septicemia                        | 2      |
|            | 038.9  | Septicemia                        | 14     |
|            | 790.7  | Bacteremia                        | 77     |
| Bone/joint | 730.02 | •                                 | 1      |
|            |        | Acute osteomyelitis pelvis        | 3      |
|            |        | Acute osteomyelitis leg           | 5      |
|            | 730.07 | Acute osteomyelitis ankle         | 2      |
|            | 730.16 | Chronic osteomyelitis leg         | 2      |
|            | 730.22 | Osteomyelitis arm                 | 2      |
|            | 730.23 | Osteomyelitis forearm             | 3      |
|            | 730.25 | Osteomyelitis pelvis              | 3      |
|            | 730.26 | Osteomyelitis leg                 | 6      |
|            | 730.27 | Osteomyelitis ankle               | 1      |
|            | 730.28 | Osteomyelitis unspecified         | 2      |
|            | 996.66 | Infection joint prosthesis        | 2      |
|            | 996.67 | Infection orthopaedic device      | 12     |
| Central    | 320.89 | Meningitis other bacteria         | 1      |
| nervous    | 320.9  | Bacterial meningitis              | 2      |
| system     | 322.9  | Meningitis unspecified            | 4      |
| Lung       | 465.9  | Acute upper respiratory infection | 3      |
| Ü          | 481    | Pneumococcal pneumonia            | 9      |
|            | 482.1  | Pseudomonas pneumonia             | 2      |
|            | 482.31 | ·                                 | 1      |
|            | 482.41 |                                   | 1      |
|            | 482.82 | E. coli pneumonia                 | 1      |
|            |        | Gram-negative pneumonia           | 10     |
|            | 482.89 | Bacterial pneumonia               | 11     |
|            | 486    | Pneumonia organism unspecified    | 35     |
|            | 510.9  | Empyema without fistula           | 10     |
|            | V46.1  | Dependence on respirator          | 12     |
| Skin/wound |        | Tracheostomy infection            | 1      |
| ,          | 528.3  | Cellulitis/abscess mouth          | 1      |
|            | 566    | Anal/rectal abscess               | 1      |
|            | 682.1  | Cellulitis/abscess neck           | 1      |
|            | 682.2  | Cellulitis trunk                  | 11     |
|            | 682.3  |                                   | 37     |
|            | 682.4  | Cellulitis hand                   | 16     |
|            | 682.5  | Cellulitis/abscess buttock        | 1      |
|            | 682.6  | Cellulitis leg                    | 77     |
|            | 682.7  | Cellulitis foot                   | 30     |
|            | 682.9  | Cellulitis/abscess unspecified    | 6      |
|            | 686.9  | Local skin infection unspecified  | 7      |
|            | 728.86 | Necrotizing fasciitis             | 1      |
|            | 785.4  | Gangrene                          | 5      |
|            | 912.1  | Abrasion shoulder/arm infected    | 5<br>1 |
|            |        |                                   |        |
|            | 916.1  | Abrasion hip/leg infected         | 3      |
|            | 916.3  | Blister hip/leg infected          | 2      |
|            | 917.3  | Blister foot and toes-infected    | 2      |
|            | 958.3  | Post-trauma wound infection       | 35     |
|            | 997.62 | Infection amputation stump        | 9      |

*Volume 66 ● Number 4* **S141** 

**Table 4** Infection ICD-9 Coding by Anatomical/Clinical Syndrome Based (continued)

| Syndrome | Code   | Code Description  | Number |
|----------|--------|---|--------|
| Other    | 041.89 | Infection bacteria other  | 39     |
|          | 112.1  | Vulva/vaginal candidiasis   | 1      |
|          | 381.4  | Nonsuppurative otitis media   | 1      |
|          | 451.82 | Superficial phlebitis arm   | 2      |
|          | 451.83 | Deep phlebitis arm  | 1      |
|          | 451.89 | Thrombophlebitis other  | 6      |
|          | 451.9  | Thrombophlebitis not otherwise stated                                   | 3      |
|          | 519.2  | Mediastinitis   | 4      |
|          | 910.5  | Infection bite head   | 5      |
|          | 916.5  | Infection bite hip/leg  | 19     |
|          | 995.91 | Systemic inflammatory response<br>syndrome without organ<br>dysfunction | 12     |
|          | 995.92 | Systemic inflammatory response syndrome with organ dysfunction          | 12     |
|          | 996.62 | Infection due to vascular device  | 18     |
|          | 996.69 | Infection due to device   | 3      |
|          | 998.59 | Other postoperative infection   | 55     |

gram-negative. 11 Because of rapid transport out of theater, most IC occurred at levels IV and V medical care.

As in previous conflicts, extremity injuries have been the most commonly encountered injury pattern. <sup>12,13</sup> During the Vietnam War, an assessment of ICs near the time of injury reported 7,106 upper extremity injuries of which 3.69% became infected and 8,838 lower extremity injuries of which 5.04% became infected. <sup>4</sup> Studies of US casualties assessing IC of extremity injuries have described rates of deep tissue or bone infections of 2% to 15%. <sup>6,9</sup> In phase II of this study, 198 of the 574 (34%) patients with extremity injuries had associated coding consistent with an IC. Although this is insightful information, the current ICD-9 codes do not specifically capture some bacteria, including *Acinetobacter*, and the more specific details on injury pattern and management strategies are not detailed. ICD-9 lacks the descriptive specificity to understand the clinical significance of these disparate rates.

It has been noted that during OIF or OEF there has been a lower proportion of thoracic wounds than in past conflicts although of those who suffer a torso injury in this study there is a relatively high rate of IC. 13 In the previously mentioned Vietnam study of wounds and infection, there were 3,490 patients with thoracic wounds with 3.84% developing an infection, and of the 2,454 patients with abdominal wounds 6.89% became infected.<sup>4</sup> An assessment of primarily non-US casualties treated on the US naval ship during the early stages of OIF revealed that abdominal injuries had an odds ratio of 2.7 for developing an infection.<sup>14</sup> A study performed in a Combat Support Hospital between September 2003 and December 2004 in Iraq revealed of the 175 (5% of the study population) primarily non-US patients with colon and rectal injuries, sepsis developed in 27 patients (16%) and had significant impact upon mortality.15

**Table 5** Infectious ICD-9 Coding by Pathogen

| Pathogen        | Code   | Code Description                               | Numb |
|-----------------|--------|--|------|
| Fungus          | 112.1  | Vulva/vaginal candidiasis                      | 1    |
|                 | 112.3  | Candidiasis of skin/nails                      | 1    |
|                 | 112.5  | Disseminated candidiasis                       | 3    |
|                 | 112.89 | Candidiasis site not available                 | 6    |
|                 | 112.9  | Candidiasis site unspecified                   | 13   |
|                 | 117.3  | Aspergillus                                    | 5    |
|                 | 117.9  | Mycoses  | 14   |
| Gram-negative   | 003.8  | Other Salmonella infection                     | 1    |
|                 | 038.42 | E. coli septicemia                             | 1    |
|                 | 038.49 | Gram-negative septicemia                       | 1    |
|                 | 041.3  | Klebsiella infection                           | 84   |
|                 | 041.4  | E. coli infection unspecified                  | 47   |
|                 | 041.6  | Proteus infection unspecified                  | 3    |
|                 | 041.7  | Pseudomonas infection                          | 102  |
|                 | 041.85 | Infection gram-negative                        | 190  |
|                 | 482.1  | Pseudomonas pneumonia                          | 3    |
| Gram-positive   | 038.11 | Staphylococcus aureus septicemia               | 6    |
|                 | 038.19 | •  | 2    |
|                 | 041.01 | , ,  | 5    |
|                 | 041.02 |  | 2    |
|                 |        | Infection Streptococcus group C                | 1    |
|                 | 041.04 |  | 39   |
|                 | 041.09 | 0 1  | 7    |
|                 | 041.11 | Infection Staphylococcus aureus                | 70   |
|                 | 041.19 |  | 27   |
|                 | 481    | Pneumococcal pneumonia                         | 2    |
| Other bacteria  | 008.45 | Clostridium difficile enteritis                | 8    |
| otrici baotoria | 032.89 |  | 1    |
|                 |        | Bacteroides fragilis infection                 | 2    |
|                 | 041.83 | <u> </u>                                       | 1    |
|                 | 041.84 | , 0  | 9    |
|                 |        | Infection bacteria other                       | 92   |
|                 | 041.9  | Bacterial infection not specified              | 29   |
|                 | 795.39 | Other nonspecific positive culture             | 11   |
|                 | V09.0  | Infection microorganism resistant penicillins  | 31   |
|                 | V09.1  | Cephalosporin resistant organism               | 1    |
|                 | V09.71 | Antimycobacterial multidrug resistant organism | 2    |
|                 | V09.80 | Other nonmultidrug resistant microorganism     | 1    |
|                 | V09.90 | Unspecified nonmultidrug resistant organism    | 1    |
|                 | V09.91 | Unspecified multidrug resistant microorganism  | 4    |

On multivariate analysis, risk factors associated with infections in this study included injuries occurring during phase II, certain mechanisms of injury (explosive device, bomb, and landmine), and higher ISS value. An analysis of casualties on the US navy ship revealed higher rates of infections with soft tissue and abdominal injuries and increasing ISS on multivariate analysis. <sup>12</sup> Although, ISS was not associated with osteomyelitis in another study of OIF or OEF casualties. <sup>9</sup> A review of a US trauma center's registry noted

S142 April Supplement 2009

Table 6 Infection Coding Based on Injury Pattern and Phase\*

|  | Phase I               |                            |                           |                   |    | Phase II              |                            |                        |                    |                       |
|--|-----------------------|----------------------------|---------------------------|-------------------|----|-----------------------|----------------------------|------------------------|--------------------|-----------------------|
|  | Extremities (n = 340) | Head and Neck<br>(n = 129) | Spine or Back<br>(n = 38) | Torso<br>(n = 87) |    | Extremities (n = 574) | Head and Neck<br>(n = 281) | Spine or Back (n = 61) | Torso<br>(n = 277) | Unspecified (n = 444) |
| Clinical and anatomical syndrome or pathogen | 27                    | 4                          | 1                         | 11                | 18 | 558                   | 268                        | 60                     | 270                | 429                   |
| Infectious complication                      | 6                     | 1                          | 0                         | 4                 | 5  | 198                   | 123                        | 33                     | 120                | 157                   |

<sup>\*</sup> Five hundred twenty-one individuals with >1 wound site on record are included in >1 column for each row.

 Table 7
 Level of Medical Care at Which Infectious Disease Diagnosis was Coded

|                                  | Level I | Level IIa | Level IIb | Level III | Level IV | Level V | Total |
|----------------------------------|---------|-----------|-----------|-----------|----------|---------|-------|
| Anatomical and clinical syndrome | 3       | 7         | 8         | 77        | 95       | 259     | 449   |
| Pathogen only                    |         |           |           | 8         | 127      | 333     | 477   |
| Infectious complication          |         |           |           | 8         | 130      | 339     | 477   |

See Ref. 10 for definitions of level of care.

**Table 8** Univariate and Multivariate Analysis of Risk Factors Associated With Development of an Infectious Complication

|  |            | Univariate              | Multivariate |            |                         |        |  |
|--|------------|-------------------------|--------------|------------|-------------------------|--------|--|
|  | Odds Ratio | 95% Confidence Interval | р            | Odds Ratio | 95% Confidence Interval | р      |  |
| Body region*   |            |                         |              |            |                         |        |  |
| Extremity  | 1.01       | 0.7–1.5                 | 0.94         |            |                         |        |  |
| Head and neck  | 1.77       | 1.32-2.38               | < 0.01       |            |                         |        |  |
| Spine and back   | 1.39       | 0.87-2.21               | 0.17         |            |                         |        |  |
| Torso  | 2.31       | 1.71-3.12               | < 0.01       |            |                         |        |  |
| Battle injury status   | 4.24       | 2.94-6.1                | < 0.01       |            |                         |        |  |
| ISS score (unit increase)  | 1.05       | 1.04-1.06               | < 0.01       |            |                         |        |  |
| ISS score 15–29  | 5.24       | 3.64-7.54               | < 0.01       | 2.33       | 1.56-3.50               | < 0.01 |  |
| ISS score >29  | 8.56       | 5.82-12.59              | < 0.01       | 3.76       | 2.47-5.73               | < 0.01 |  |
| Phase II   | 38.1       | 16.75-86.46             | < 0.01       | 20.25      | 8.58-47.82              | < 0.01 |  |
| Mechanism of injury <sup>†</sup>   |            |                         |              |            |                         |        |  |
| Explosive device   | 6.89       | 4.08-11.63              | < 0.01       | 1.86       | 1.04-3.33               | 0.04   |  |
| Bomb   | 8.13       | 2.27-29.13              | < 0.01       | 9.97       | 1.81-54.82              | < 0.01 |  |
| Grenade  | 2.34       | 1.21-4.52               | 0.01         |            |                         |        |  |
| Gunshot  | 2.71       | 1.49-4.94               | 0.01         |            |                         |        |  |
| Landmine   | 7.51       | 2.91-19.38              | < 0.01       | 4.0        | 1.29-12.4               | 0.02   |  |
| Mortar   | 3.60       | 1.66-7.78               | < 0.01       |            |                         |        |  |
| Unexploded ordnance  | 1.63       | 0.34-7.81               | 0.54         |            |                         |        |  |
| Motor vehicle crash  | 1.48       | 0.70-3.12               | 0.31         |            |                         |        |  |
| Machinery/equipment  | 0.83       | 0.27-2.55               | 0.04         |            |                         |        |  |
| Burn, fall/jump from height,<br>knife or other sharp<br>object, Shrapnel/<br>fragment <sup>‡</sup> | 0.34       | 0.097–1.15              | 0.08         |            |                         |        |  |

<sup>\*</sup> Reference group 'Unspecified' (i.e., Barell Matrix Body Region unclassifiable by site).

certain factors to be associated with sepsis to include comorbidities, immunosuppression, Glascow Coma Scale (GCS), and increasing ISS. <sup>16</sup> Although comorbidities, such as diabetes or immunosuppression, were associated with sepsis, these are unlikely occurrences in a population of healthy deployed

military personnel. Another predictor was a GCS score of <8; however, the majority of patients suffered a blunt mechanism of injury (86%) in the civilian population. Finally, higher ISS value was associated with sepsis among civilian trauma patients as in our study.

*Volume 66* • *Number 4* **\$143** 

<sup>&</sup>lt;sup>†</sup> Reference group "Other," as implicitly defined in Table 2.

<sup>&</sup>lt;sup>‡</sup> Categories combined due to sparse data.

Although this is the first broad assessment of ICs of a purely US population of combat-related injuries, there are a number of limitations in the use of these data. The JTTR has undergone a series of improvements in data collection and entry since its conception, leading to a possibility of discrepancies in data over time which might be reflective of the differences reported between phase I and II in contrast to the differences in ISS or mechanisms or injury. Also, the collection of level V data has been incomplete at many facilities. Unfortunately, this is evident by the lack of outcome data for the majority of casualties. Fortunately, the latest version of the JTTR is being adequately staffed at the major Army level V facilities to minimize this concern. Although charts are now being entered serially into the JTTR by Medical Treatment Facility Unit, the incomplete yearly record completion rate might be associated with a chart selection bias that is not evident at this time. Although, the JTTR is designed as a DoD process, the JTTR was not being implemented at level V Navy and Air Force facilities at the time of this review but is now occurring. There is also a limitation of ICD-9 diagnosis for ICs as they do not provide the clinical granularity necessary to impact patient care. To address this issue, the latest version of the JTTR has a specific infectious disease module being added to enhance infectious disease specific data collection. There are also other performance improvement and research programs under development that could augment the JTTR. This includes the development of a DoD Joint Bacterial Repository to collect infection specific data with associated bacterial isolates. In addition, the Infectious Disease Clinical Research Program is developing a Trauma Infectious Disease Outcome Study for long-term follow-up of casualties to determine chronic combat-related injury ICs.

This initial review of the JTTR has provided some unique descriptions of the challenges facing healthcare providers to mitigate the acute and chronic ICs of combat casualties. There are a number of limitations of the current registry that are actively being addressed. Despite these limitations, data collected from the JTTR will enable improvements in ongoing data collection and provide insight into further improvements in patient care and future research programs. Infectious events occur frequently in combat casualties and likely cause significant adverse clinical outcomes. Understanding the ep-

idemiology is the first step toward an evidence-based intervention program.

## **REFERENCES**

- Eastridge BJ, Jenkins D, Flaherty S, Schiller H, Holcomb JB.
   Trauma system development in a theater of war: experiences from
   Operation Iraqi Freedom and Operation Enduring Freedom.
   *J Trauma*. 2006;61:1366–1372; discussion 1372–1373.
- Arnold K, Cutting RT. Causes of death in United States Military personnel hospitalized in Vietnam. Mil Med. 1978;143:161–164.
- Feltis JJ. Surgical experience in a combat zone. Am J Surg. 1970; 119:275–278.
- Hardaway RM III. Viet Nam wound analysis. J Trauma. 1978; 18:635–643.
- Kelly JF, Ritenour AE, McLaughlin DF, et al. Injury severity and causes of death from Operation Iraqi Freedom and Operation Enduring Freedom: 2003–2004 versus 2006. *J Trauma*. 2008;64(2 Suppl):S21–S26; discussion S26–S27.
- Lin DL, Kirk KL, Murphy KP, McHale KA, Doukas WC. Evaluation of orthopaedic injuries in Operation Enduring Freedom. *J Orthop Trauma*. 2004;18:300–305.
- Davis KA, Moran KA, McAllister CK, Gray PJ. Multidrug-resistant Acinetobacter extremity infections in soldiers. Emerg Infect Dis. 2005;11:1218–1224.
- Johnson EN, Burns TC, Hayda RA, Hospenthal DR, Murray CK. Infectious complications of open type III tibial fractures among combat casualties. *Clin Infect Dis.* 2007;45:409–415.
- Yun HC, Branstetter JG, Murray CK. Osteomyelitis in military personnel wounded in Iraq and Afghanistan. *J Trauma*. 2008;64(2 Suppl):S163–S168; discussion S168.
- Murray CK. Epidemiology of infections associated with combatrelated injuries in Iraq and Afghanistan. *J Trauma*. 2008;64(3 Suppl):S232–S238.
- Murray CK, Hinkle MK, Yun HC. History of infections associated with combat-related injuries. *J Trauma*. 2008;64(3 Suppl):S221– S231.
- Owens BD, Kragh JF Jr, Macaitis J, Svoboda SJ, Wenke JC. Characterization of extremity wounds in Operation Iraqi Freedom and Operation Enduring Freedom. J Orthop Trauma. 2007;21:254– 257
- Petersen K, Riddle MS, Danko JR, et al. Trauma-related infections in battlefield casualties from Iraq. Ann Surg. 2007;245:803–811.
- Owens BD, Kragh JF, Wenke JC, Macaitis J, Wade CE, Holcomb JB. Combat wounds in Operation Iraqi Freedom and Operation Enduring Freedom. *J Trauma*. 2008;64:295–299.
- Steele SR, Wolcott KE, Mullenix PS, et al. Colon and rectal injuries during Operation Iraqi Freedom: are there any changing trends in management or outcome? *Dis Colon Rectum.* 2007;50:870–877.
- Osborn TM, Tracy JK, Dunne JR, Pasquale M, Napolitano LM. Epidemiology of sepsis in patients with traumatic injury. *Crit Care Med.* 2004;32:2234–2240.

S144 April Supplement 2009